

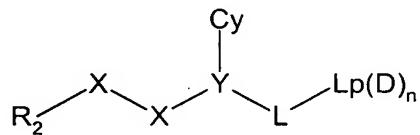
Continuation of Serial No. 10/030,187

Amendments to the Claims:

This listing of claims will replace all prior versions, and listings, of claims in the application.

Listing of Claims:

1 (currently amended) : A serine protease inhibitor compound of formula (I)



(I)

wherein:

R<sub>2</sub> is:-

(i) phenyl optionally being substituted in the 3 and/or 4 position by halo, nitro, thiol, haloalkoxy, hydrazido, alkylhydrazido, amino, cyano, haloalkyl, alkylthio, alkenyl, alkynyl, acylamino, tri or difluoromethoxy, carboxy, acyloxy, MeSO<sub>2</sub>- or R<sub>1</sub>, and optionally substituted at the 6 position by amino, hydroxy, halo, alkyl, carboxy, alkoxycarbonyl, cyano, amido, aminoalkyl, alkoxy or alkylthio;

(ii) naphth-2-yl optionally substituted at the 6 or 7 position by halo, haloalkoxy, haloalkyl, cyano, nitro, amino, hydrazido, alkylthio, alkenyl, alkynyl or R<sub>1j</sub> and optionally substituted at the 3 position by amino, hydroxy, halo, alkyl, carboxy, cyano, amido, aminoalkyl, alkoxy or alkylthio;

(iii) isoquinolin-7-yl, indol-5-yl, indol-6-yl, indazol-5-yl, indazol-6-yl, benzothiazol-6-yl or benzisoxazol-5-yl optionally substituted at the 3 position by halo, haloalkoxy, haloalkyl, cyano, nitro, amino, hydrazido, alkylthio, alkenyl, alkynyl or R<sub>1j</sub>;

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(iv) benzimidazol-5-yl or benzothiazol-6-yl optionally substituted at the 2 position by amino;

(v) thien-2-yl or thien-3-yl optionally substituted at the 4 or 5 position by halo, haloalkoxy, haloalkyl, cyano, nitro, amino, hydrazido, alkylthio, alkenyl, alkynyl or  $R_1$ ;

(vi) 3,4-methylenedioxophenyl, 2,3-dihydroindol-6-yl, 3,3-dichloro-2-oxo-indol-6-yl or 1-methyl-3-aminoindazol-5-yl;

(vii) benzothiazol-2-yl, imidazo[1,2-a]pyrimidin-2-yl or tetrahydroimidazo[1,2-a]pyrimidin-2-yl;

(viii) pyrazol-2-yl optionally substituted at the 5 position by halo, haloalkoxy, haloalkyl, cyano, nitro, amino, hydrazido, alkylthio, alkenyl, alkynyl or  $R_1$ ;

(ix) pyrid-2-yl optionally substituted at the 5 position by halo, haloalkoxy, haloalkyl, cyano, nitro, amino, hydrazido, alkylthio, alkenyl, alkynyl or  $R_1$ ;

(x) pyrid-3-yl optionally substituted at the 6 position by halo, haloalkoxy, haloalkyl, cyano, nitro, amino, hydrazido, alkylthio, alkenyl, alkynyl or  $R_1$ ;

(xi) benzofur-2-yl optionally substituted at the 3 position by amino, hydroxy, halo, alkyl, carboxy, cyano, amido, aminoalkyl, alkoxy or alkylthio and at the 5 or 6 position by halo, haloalkoxy, haloalkyl, cyano, nitro, amino, hydrazido, alkylthio, alkenyl, alkynyl or  $R_{1j}$ ;

(xii) indol-2-yl optionally substituted on the indole nitrogen atom by alkyl and optionally substituted at the 5 or 6 position by halo, haloalkoxy, haloalkyl, cyano, nitro, amino, hydrazido, alkylthio, alkenyl, alkynyl or  $R_{1j}$ ;

(xiii) indol-6-yl substituted at the 5 position by amino, hydroxy, halo, alkyl, carboxy, alkoxycarbonyl, cyano, amido, aminoalkyl, alkoxy or alkylthio and optionally substituted at the 3 position by halo, haloalkoxy, haloalkyl, cyano, nitro, amino, hydrazido, alkylthio, alkenyl, alkynyl or  $R_{1j}$ ; or

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(xiv) benzo [b] thiophen-2-yl optionally substituted at the 3 position by amino, hydroxy, halo, alkyl, carboxy, cyano, amido, aminoalkyl, alkoxy or alkylthio and at the 5 or 6 position by halo, haloalkoxy, haloalkyl, cyano, nitro, amino, hydrazido, alkylthio, alkenyl, alkynyl or R<sub>1j</sub>; a 5 or 6 membered aromatic carbon ring optionally interrupted by a nitrogen, oxygen or sulphur ring atom, optionally being substituted in the 3 and/or 4 position (in relation to the point of attachment of X-X) by halo, nitro, thiol, haloalkoxy, hydrazido, alkylhydrazido, amino, cyano, haloalkyl, alkylthio, alkenyl, alkynyl, acylamino, tri or difluoromethoxy, carboxy, aeyloxy, MeSO<sub>2</sub> or R<sub>1</sub>, or the substituents at the 3 and 4 positions taken together form a fused ring which is a 5 or 6 membered carbocyclic or heterocyclic ring optionally substituted by halo, haloalkoxy, haloalkyl, cyano, nitro, amino, hydrazido, alkylthio, alkenyl, alkynyl or R<sub>1j</sub>, and optionally substituted in the position alpha to the X-X group (i.e. 6 position for a six membered aromatic ring etc) by amino, hydroxy, halo, alkyl, carboxy, alkoxycarbonyl, cyano, amido, aminoalkyl, alkoxy or alkylthio

with the proviso that R<sub>2</sub> cannot be aminoisoquinolyl;

-X-X- is -CONH- each X independently is a C, N, O or S atom or a CO, CR<sub>1a</sub>, C(R<sub>1a</sub>)<sub>2</sub> or NR<sub>1a</sub> group, at least one X being C, CO, CR<sub>1a</sub> or C(R<sub>1a</sub>)<sub>2</sub>;

each R<sub>1a</sub> independently represents hydrogen or hydroxyl, alkoxy, alkyl, aminoalkyl, hydroxyalkyl alkoxylalkyl, alkoxycarbonyl, alkylaminocarbonyl, alkoxycarbonylamine, aeyloxymethoxy carbonyl or alkylamine optionally substituted by hydroxy, alkylamine, alkoxy, o xo, aryl or cycloalkyl,

R<sub>1</sub> is hydrogen, hydroxy, alkoxy, alkyl, alkylaminoalkyl, alkanoyl, hydroxyalkyl, alkoxyalkyl, alkoxycarbonyl, alkylaminocarbonyl, alkylamino, carboxyl, carboxymethyl, amido or amidomethyl;

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R<sub>1j</sub> is hydrogen, hydroxy, alkoxy, alkyl, alkanoyl, hydroxyalkyl, alkoxyalkyl, alkoxycarbonyl, alkylamino, carboxyl, carboxymethyl, amido or amidomethyl; as defined for R<sub>1a</sub>, provided that R<sub>1</sub> is not unsubstituted aminealkyl;

Y (the α-atom) is a CH nitrogen atom or a CR<sub>1b</sub> group;

Cy is an optionally R<sub>3a</sub> substituted: phenyl, pyridyl, thienyl, thiazolyl, naphthyl, piperidinyl, furanyl, pyrrolyl, isoxazolyl, isothiazolyl, pyrazolyl, oxazolyl, imidazolyl, 1,2,4-thiadiazolyl, 1,3,4-thiadiazolyl, pyrimidinyl, pyridazinyl, quinolyl, isoquinolyl, benzofuryl, benzothienyl or cycloalkyl group, or a phenyl group substituted by R<sub>3i</sub>X<sub>i</sub>; a saturated or unsaturated, mono or poly cyclic, homo or heterocyclic group, optionally substituted by groups R<sub>3a</sub> or R<sub>3i</sub>X<sub>i</sub>;

each R<sub>3a</sub> independently is hydrogen, hydroxyl, alkoxy, aralkyloxy, alkyl, alkylaminoalkyl, hydroxymethyl, carboxy, alkoxyalkyl, alkoxycarbonyl, alkylaminocarbonyl, aminomethyl, CONH<sub>2</sub>, CH<sub>2</sub>CONH<sub>2</sub>, (1-6C) alkanoylamino, alkoxycarbonylaminoR<sub>1e</sub>, amino, halo, cyano, nitro, thiol, alkylthio, alkylsulphonyl, alkylsulphenyl, triazolyl, imidazolyl, tetrazolyl, hydrazido, alkylimidazolyl, thiazolyl, alkylthiazolyl, alkylloxazolyl, oxazolyl, alkylsulphonamido, alkylaminosulphonyl, aminosulphonyl, haloalkoxy, haloalkyl, a group of the formula -C(X<sup>3</sup>)N(R<sup>11</sup>)R<sup>12</sup> (wherein X<sup>3</sup> is O or S; and R<sup>11</sup> and R<sup>12</sup> are independently selected from hydrogen, methyl or ethyl or together with the nitrogen atom to which they are attached form a pyrrolidin-1-yl, piperidin-1-yl or morpholino group), or -OCH<sub>2</sub>O- which is bonded to two adjacent ring atoms in Cy;

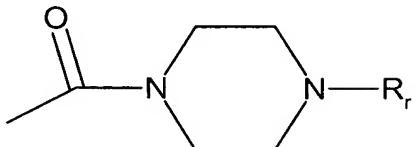
X<sub>i</sub> is a bond, O, NH or CH<sub>2</sub>;

R<sub>3i</sub> is phenyl, pyridyl or pyrimidinyl optionally substituted by R<sub>3a</sub>;

R<sub>1b</sub>, R<sub>1e</sub> and R<sub>1j</sub> are as defined for R<sub>1a</sub>; and

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-L-Lp(D)<sub>n</sub> is of the formula:

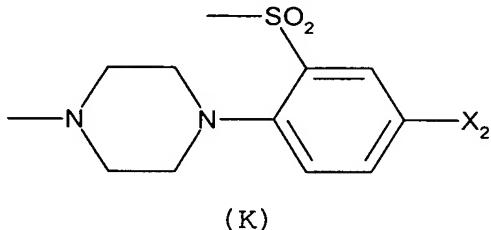


in which R<sub>r</sub> is -(CH<sub>2</sub>)<sub>c</sub>-R<sub>C</sub>, -CHR<sub>e</sub>R<sub>f</sub>, -CH<sub>2</sub>-CHR<sub>e</sub>R<sub>f</sub>, -CH<sub>2</sub>-CH<sub>2</sub>-CHR<sub>e</sub>R<sub>f</sub>, or R<sub>g</sub> in which c is 1 or 2; R<sub>C</sub> is thienyl, thiazolyl (which may bear an amino substituent), isothiazolyl, oxazolyl, isoxazolyl, pyrazolyl, imidazolyl, pyridyl (which may bear an alkylsulphonyl, aminosulphonyl, alkylaminosulphonyl, alkylaminocarbonyl, amino, amido, (1-4C)alkoxycarbonyl, carboxy, acetyl amino, chloro, fluoro, cyano, (1-3C)alkyl, trifluoromethyl, methoxy, ethoxy, nitro, hydroxy, alkylsulphonylamino, triazolyl or tetrazolyl substituent), pyrimidinyl, pyridazinyl, pyrazinyl or phenyl (which may bear a methyl, methylamino, dimethylamino, carboxy, dialkylaminosulphonyl, alkylsulphonyl, aminosulphonyl, alkylaminosulphonyl, alkylaminocarbonyl, amino, amido, alkoxy carbonyl, acetyl amino, chloro, fluoro, cyano, methoxy, ethoxy, nitro, hydroxy, alkylsulphonylamino, triazolyl or tetrazolyl substituent); each of R<sub>e</sub> and R<sub>f</sub> independently is hydrogen or C<sub>1-3</sub>alkyl; or CHR<sub>e</sub>R<sub>f</sub> is cyclopentyl (which may bear a hydroxy, amino, (1-3C)alkoxy, (1-3C)hydroxyalkyl, (1-3C)alkyl, carboxy, methoxycarbonyl or ethoxycarbonyl substituent at the 3- or 4-position), cyclohexyl (which may bear a hydroxy, amino, (1-3C)alkoxy, (1-3C)hydroxyalkyl, (1-3C)alkyl, carboxy, methoxycarbonyl or ethoxycarbonyl substituent at the 3- or 4-position), tetrahydropyran-4-yl, tetrahydrothiopyran-4-yl, pyrrolidin-3-yl (which may bear a hydroxy, amino, (1-3C)alkoxy, (1-3C)hydroxyalkyl, (1-3C)alkyl, carboxy, methoxycarbonyl or ethoxycarbonyl substituent at the 1-position), piperidin-4-yl (which may bear a hydroxy, amino, (1-3C)alkoxy, (1-3C)hydroxyalkyl, (1-3C)alkyl, carboxy, methoxycarbonyl or ethoxycarbonyl substituent at the 1-

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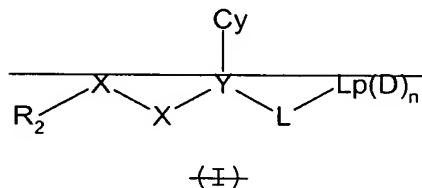
position), or indan-2-yl; and R<sub>g</sub> is 2-methylsulphonylphenyl which may bear a 4-fluoro substituent or R<sub>g</sub> is λ<sup>6</sup>-1,1-dioxobenzo[b]thiophen-7-yl;

or a physiologically-tolerable salt thereof;  
provided that Lp(D)<sub>n</sub> is not of the formula (K) :



wherein X<sub>2</sub> is fluoro or hydrogen.

2 (currently amended) : A compound according to claim 1  
~~serine protease inhibitor compound of formula (I)~~



wherein:

R<sub>2</sub> is a 5 or 6 membered aromatic carbon ring optionally interrupted by a nitrogen, oxygen or sulphur ring atom, optionally being substituted in the 3 and/or 4 position (in relation to the point of attachment of X-X) by halo, nitro, thiol, haloalkoxy, hydrazido, alkylhydrazido, amino, cyano, haloalkyl, alkylthio, alkenyl, alkynyl, acylamino, tri or difluoromethoxy, carboxy, acyloxy, MesO<sub>2</sub> or R<sub>1</sub>, or the substituents at the 3 and 4 positions taken together form a fused ring which is a 5 or 6 membered carbocyclic or heterocyclic ring optionally substituted by halo, haloalkoxy, haloalkyl, cyano, nitro, amino, hydrazido, alkylthio, alkenyl, alkynyl or R<sub>1j</sub>, and optionally substituted in the position alpha to the X-X group (i.e. 6 position for a six membered

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~~aromatic ring etc) by amino, hydroxy, halo, alkyl, carboxy, alkoxy carbonyl, cyano, amido, aminoalkyl, alkoxy or alkylthio with the proviso that R<sub>2</sub> cannot be aminoisouinolyl;~~

~~each X independently is a C, N, O or S atom or a CO, CR<sub>1a</sub>, C(R<sub>1a</sub>)<sub>2</sub> or NR<sub>1a</sub> group, at least one X being C, CO, CR<sub>1a</sub> or C(R<sub>1a</sub>)<sub>2</sub>;~~

~~each R<sub>1a</sub> independently represents hydrogen or hydroxyl, alkoxy, alkyl, aminoalkyl, hydroxyalkyl, alkoxyalkyl, alkoxy carbonyl, alkylaminocarbonyl, alkoxy carbonyl amino, acyloxymethoxy carbonyl or alkylamino optionally substituted by hydroxy, alkylamino, alkoxy, exo, aryl or cycloalkyl;~~

~~R<sub>1</sub> is as defined for R<sub>1a</sub>, provided that R<sub>1</sub> is not unsubstituted aminoalkyl;~~

~~Y (the α atom) is a nitrogen atom or a CR<sub>1b</sub> group;~~

~~Cy is an optionally R<sub>3a</sub> substituted: phenyl, pyridyl, thienyl, thiazolyl, naphthyl, piperidinyl or cycloalkyl group; a saturated or unsaturated, mono or poly cyclic, homo or heterocyclic group, optionally substituted by groups R<sub>3a</sub> or phenyl optionally substituted by R<sub>3a</sub>;~~

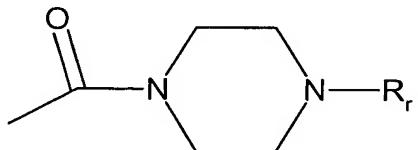
each R<sub>3a</sub> independently is hydrogen, hydroxyl, methoxy, ethoxy, methyl, ethyl, methylaminomethyl, dimethylaminomethyl, hydroxymethyl, carboxy, methoxymethyl, methoxycarbonyl, ethoxycarbonyl, methylaminocarbonyl, dimethylamino-carbonyl, aminomethyl, CONH<sub>2</sub>, CH<sub>2</sub>CONH<sub>2</sub>, acetyl amino, methoxycarbonylamino, ethoxycarbonylamino, t-butoxycarbonylamino, amino, fluoro, chloro, cyano, nitro, thiol, methylthio, methylsulphonyl, ethylsulphonyl, methylsulphenyl, methylsulphonylamido, ethylsulphonylamido, methylaminosulphonyl, ethylaminosulphonyl, aminosulphonyl, trifluoromethoxy or trifluoromethyl; R<sub>1e</sub>, amino, halo, cyano, nitro, thiol, alkylthio, alkylsulphonyl, alkylsulphenyl, triazolyl, imidazolyl, tetrazolyl, hydrazide, alkyl imidazolyl, thiazolyl, alkyl thiazolyl, alkyl oxazolyl,

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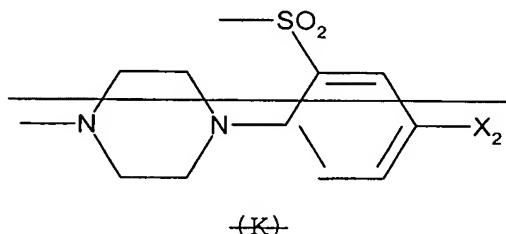
~~oxazolyl, alkylsulphonamido, alkylaminosulphonyl, aminosulphonyl, haloalkoxy and haloalkyl,~~

~~R<sub>1b</sub>, R<sub>1e</sub> and R<sub>1j</sub> are as defined for R<sub>1a</sub>, and~~

~~-L-Lp(D)<sub>n</sub> is of the formula:~~

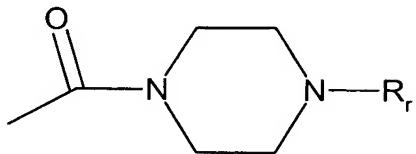


in which R<sub>r</sub> is -(CH<sub>2</sub>)<sub>c</sub>-R<sub>c</sub>, -CHR<sub>e</sub>R<sub>f</sub>, -CH<sub>2</sub>-CHR<sub>e</sub>R<sub>f</sub>, or R<sub>g</sub> in which c is 1 or 2; R<sub>c</sub> is pyridyl or phenyl (which phenyl may bear a fluoro, chloro, methyl, CONH<sub>2</sub>, SO<sub>2</sub>NH<sub>2</sub>, methylaminosulphonyl, dimethylaminosulphonyl, methoxy or methylsulphonyl substituent); each of R<sub>e</sub> and R<sub>f</sub> independently is hydrogen or C<sub>1-3</sub>alkyl; or CHR<sub>e</sub>R<sub>f</sub> is cyclopentyl (which may bear a methyl, ethyl or hydroxymethyl substituent at the 3- or 4-position), cyclohexyl (which may bear a methyl, ethyl or hydroxymethyl, (1-3C)alkyl, carboxy, methoxycarbonyl or ethoxycarbonyl substituent at the 3- or 4-position), tetrahydropyran-4-yl, tetrahydrothiopyran-4-yl, pyrrolidin-3-yl (which may bear a 1-methyl substituent), piperidin-4-yl (which may bear a 1-methyl substituent) or indan-2-yl; and R<sub>g</sub> is 2-methylsulphonylphenyl which may bear a 4-fluoro substituent or R<sub>g</sub> is λ<sup>6</sup>-1,1-dioxobenzo[b]thiophen-7-yl, ~~or a physiologically tolerable salt thereof,~~ provided that Lp(D)<sub>n</sub> is not of the formula (K) →



wherein X<sub>2</sub> is fluoro or hydrogen.

3 (original): A compound according to claim 1 wherein -L-Lp(D)<sub>n</sub> is of the formula:



in which R<sub>r</sub> is -(CH<sub>2</sub>)<sub>c</sub>-R<sub>c</sub>; in which c is 2; R<sub>c</sub> is thiaryl, thiazolyl (which may bear an amino substituent), isothiazolyl, oxazolyl, isoxazolyl, pyrazolyl, imidazolyl, pyridyl (which may bear an amino, methoxycarbonyl, carboxy, fluoro, cyano, methyl, methylsulphonyl, aminosulphonyl, methylaminosulfonyl, dimethylaminosulfonyl, or trifluoromethyl substituent), pyrimidinyl, pyridazinyl, pyrazinyl or phenyl (which phenyl may bear a fluoro, chloro, cyano, methyl, amino, methylsulphonyl, aminosulphonyl, methylaminosulphonyl, dimethylaminosulphonyl, methylamino, dimethylamino, carboxy, methoxycarbonyl or methoxy substituent).

4 (currently amended): A compound according to claim any one of claims 1 to 3 wherein R<sub>c</sub> is thiazolyl (which may bear an amino substituent), pyrimidinyl, pyrazolyl, imidazolyl, pyridyl (which may bear a methylsulphonyl, aminosulphonyl, methylaminosulfonyl, dimethylaminosulfonyl, fluoro, cyano, methyl or trifluoromethyl substituent), pyridazinyl, pyrazinyl or phenyl (which phenyl may bear a fluoro, chloro, cyano, methyl, amino, methylamino, dimethylamino, carboxy, methoxycarbonyl, methylsulphonyl, aminosulphonyl, methylaminosulfonyl, dimethylaminosulfonyl, or methoxy substituent).

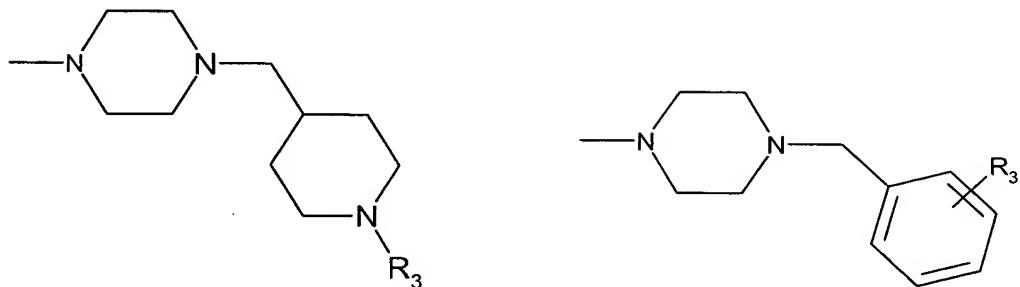
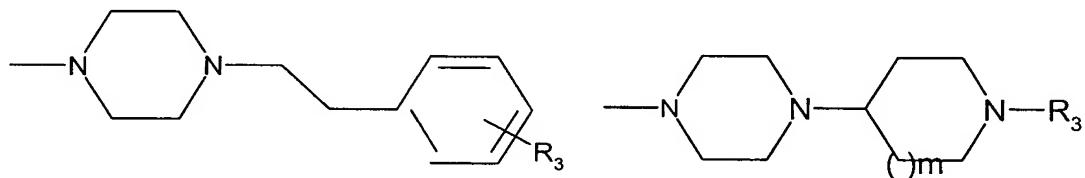
5 (currently amended): A compound according to claim any one of claims 1 to 4 wherein R<sub>c</sub> is thiazolyl (which may bear an amino substituent), pyrazolyl, imidazolyl, pyridyl (which may

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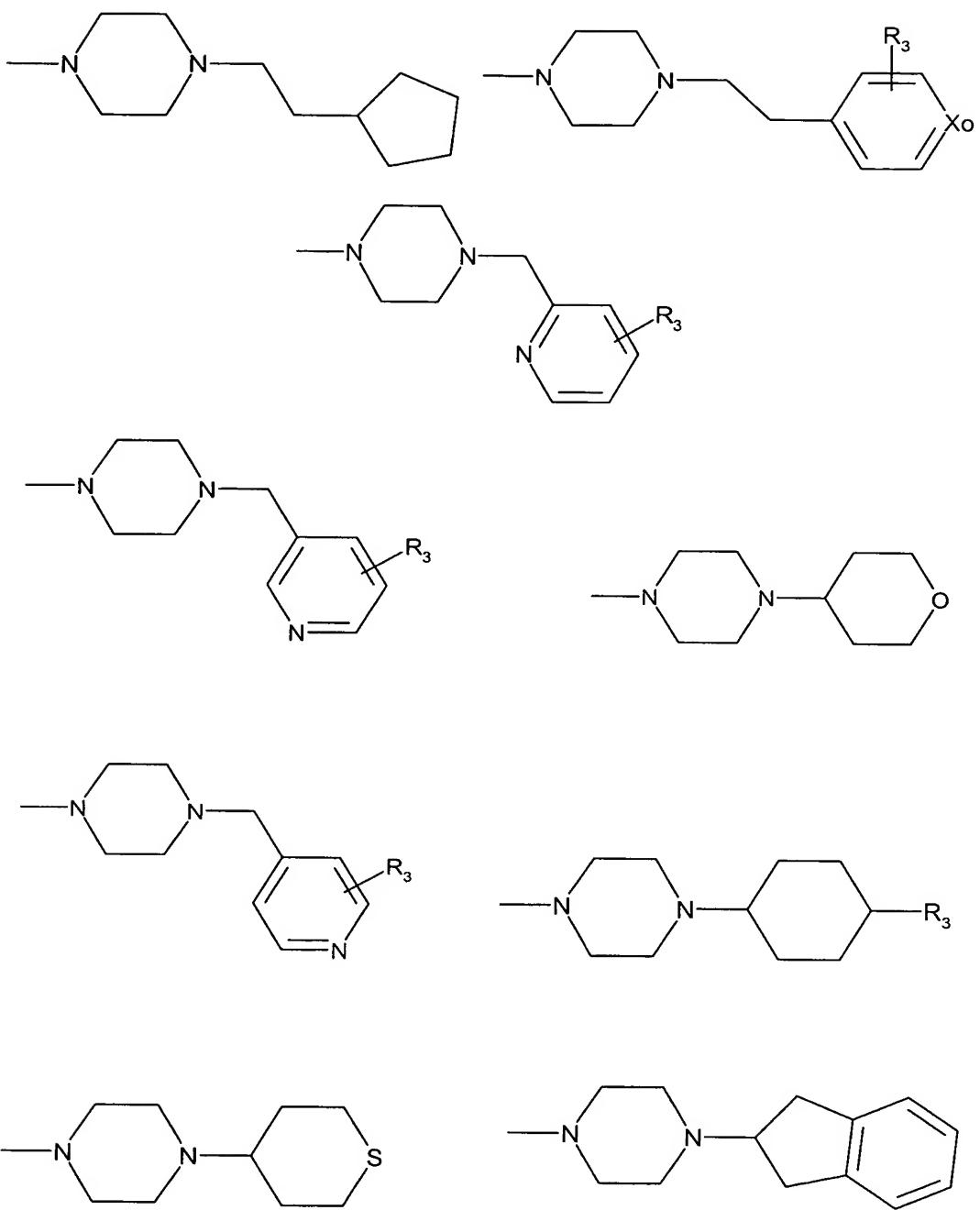
bear a fluoro, cyano, methyl or trifluoromethyl substituent), pyridazinyl or pyrazinyl.

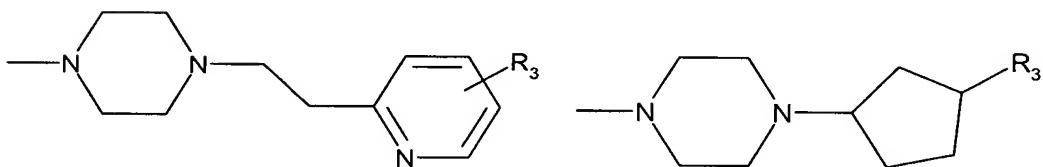
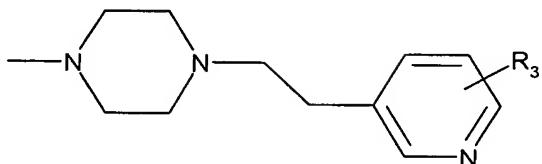
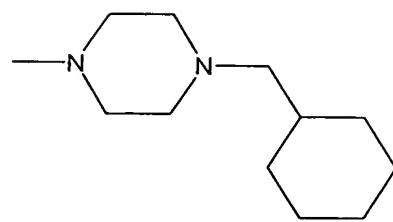
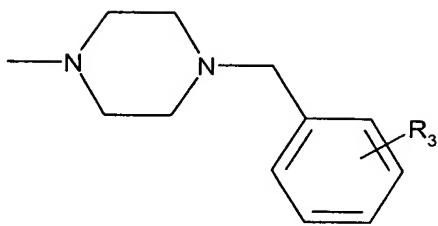
6 (currently amended): A compound according to claim any one of claims 1 to 5 wherein Rc is thiazol-2-yl, 2-aminothiazol-4-yl, pyrazol-1-yl, pyrazol-4-yl, pyridazin-3-yl, imidazol-1-yl, imidazol-4-yl, pyrazin-2-yl, pyrid-2-yl, pyrid-3-yl, pyrid-4-yl, 3-fluoropyrid-4-yl, 2-cyanopyrid-4-yl, 2-methylpyrid-4-yl or 2-trifluoromethylpyrid-6-yl.

7 (currently amended): A compound according to claim 1 wherein L is CO and -Lp(D)n is of the formula:



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wherein;

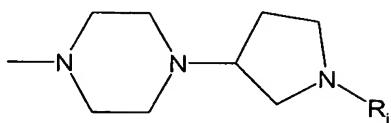
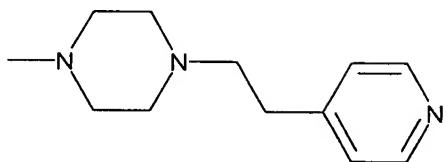
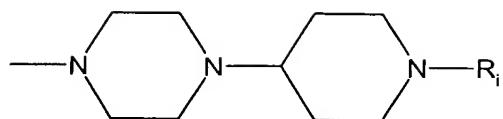
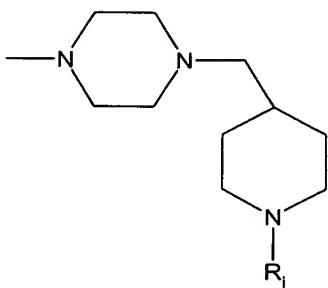
m represents 0 or 1;

x<sup>0</sup> represents CH or N; and

when R<sub>3</sub> is present as a substituent on an aromatic ring, it is selected from hydrogen, alkylsulphonyl, aminosulphonyl, alkylaminosulphonyl, alkylaminocarbonyl, amino, amido, alkoxy carbonyl, acetyl amino, chloro, fluoro, cyano, methoxy, ethoxy, nitro, hydroxy, alkylsulphonylamino, triazolyl and tetrazolyl; and

when R<sub>3</sub> is present as a substituent on a saturated ring, it is selected from hydrogen, hydroxy, amino, (1-3C) alkoxy, (1-3C) hydroxyalkyl, (1-3C) alkyl, carboxy, methoxycarbonyl and ethoxycarbonyl.

8 (original): A compound according to claim 7 wherein -Lp(D)<sub>n</sub> is of the formula:



wherein R<sub>1</sub> is hydrogen or (1-6C)alkyl.

9 (currently amended): A compound according to claim any one of claims 1 to 8 wherein R<sub>2</sub> is: - phenyl, thien-2-yl, naphthyl, indol-2-yl, indol-6-yl, benzo[b]furan-5-yl, benzo[b]thiophen-2-yl or benzimidazol-2-yl (each of which is optionally substituted as defined in claim 1).

(i) phenyl optionally being substituted in the 3 and/or 4 position by fluoro, chloro, bromo, iodo, nitro, difluoromethoxy, trifluoromethoxy, amino, cyano, trifluoromethyl, methylthio, vinyl, carboxy, acetoxy, MeSO<sub>2</sub>-, hydroxy, methoxy, ethoxy, methyl, methoxycarbonyl, methylamino, ethylamino or amido, and optionally substituted at the 6 position by amino, hydroxy, fluoro, methoxycarbonyl, cyano or aminomethyl;

(ii) naphth-2-yl optionally substituted at the 6, position by hydroxy and optionally substituted at the 3 position by amino or hydroxy;

(iii) isoquinolin-7-yl, indol-5-yl, indol-6-yl, indazol-5-yl, indazol-6-yl, benzothiazol-6-yl or benzisoxazol-5-yl optionally substituted at the 3 position by chloro, bromo, amino, methyl or methoxy;

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- (iv) benzimidazol-5-yl or benzothiazol-6-yl optionally substituted at the 2 position by amino;
- (v) thien-2-yl or thien-3-yl optionally substituted at the 4 or 5 position by methylthio, methyl or acetyl;
- (vi) 3,4-methylenedioxypyhenyl, 2,3-dihydroindol-6-yl, 3,3-dichloro-2-oxo-indol-6-yl or 1-methyl-3-aminoindazol-5-yl;
- (vii) benzothiazol-2-yl, imidazo[1,2-a]pyrimidin-2-yl or tetrahydroimidazo[1,2-a]pyrimidin-2-yl;
- (viii) pyrazol-2-yl substituted at the 5 position by methyl;
- (ix) pyrid-2-yl optionally substituted at the 6 position by chloro;
- (x) pyrid-3-yl optionally substituted at the 4 position by chloro;
- (xi) benzofur-2-yl optionally substituted at the 3 position by chloro, methyl or methoxy, at the 5 or 6 position by methyl and at the 6 position by methoxy;
- (xii) indol-2-yl optionally substituted on the indole nitrogen atom by methyl and optionally substituted at the 5 or 6 position by fluoro, chloro, bromo, methyl or methoxy;
- (xiii) indol-6-yl substituted at the 5 position by chloro, fluoro or hydroxy and optionally substituted at the 3 position by chloro or methyl; or
- (xiv) benzo[b]thiophen-2-yl optionally substituted at the 3 position by fluoro, chloro or methyl, and optionally substituted at the 5 or 6 position by fluoro, chloro, methyl, hydroxy, or methoxy.

10 (currently amended): A compound according to claim any one of claims 1 to 9 wherein optional substituents for R<sub>2</sub> are selected from:-

- (i) phenyl, 2-aminophenyl, 3-aminophenyl, 2-amino-3-fluorophenyl, 2-amino-4-fluorophenyl, 2-amino-4-chlorophenyl,

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2-amino-3-bromophenyl, 2-amino-3-nitrophenyl, 2-amino-4-nitrophenyl, 3,4-dimethoxy-5-aminophenyl, 2-amino-4-methylphenyl, 2-amino-3-methylphenyl, 2-amino-3-methoxyphenyl, 3,4-diaminophenyl, 3,5-diaminophenyl, 3-amino-4-fluorophenyl, 3-amino-4-chlorophenyl, 3-amino-4-bromophenyl, 3-amino-4-hydroxyphenyl, 3-amino-4-carboxymethylphenyl, 3-amino-4-methylphenyl, 3-amino-4-methoxyphenyl, 2-fluorophenyl, 4-fluoro-3-cyanophenyl, 3-chlorophenyl, 3-chloro-4-hydroxphenyl, 3-chloro-5-hydroxyphenyl, 4-chlorophenyl, 4-chloro-2-hydroxyphenyl, 4-chloro-3-hydroxyphenyl, 4-chloro-3-methylphenyl, 4-chloro-3-methoxyphenyl, 4-bromophenyl, 4-bromo-3-methylphenyl, 4-iodophenyl, 2-cyanophenyl, 3-cyanophenyl, 4-cyanophenyl, 3-cyano-5-aminophenyl, 2-hydroxyphenyl, 2-hydroxy-4-methoxyphenyl, 3-hydroxphenyl, 3-hydroxy-4-methylphenyl, 2,4-dihydroxyphenyl, 3,4-dihydroxyphenyl, 3-hydroxy-4-methoxyphenyl, 4-difluoromethoxyphenyl, 4-trifluoromethoxphenyl, 4-trifluoromethylphenyl, 4-methylthiophenyl, 4-methoxycarbonylphenyl, 4-acetoxyphenyl, 4-methanesulfonylphenyl, 3-methylphenyl, 3-methyl-5-aminophenyl, 4-methylphenyl, 4-vinylphenyl, 4-methoxyphenyl, 4-ethoxyphenyl, 4-methoxy-3-chlorophenyl, 4-methoxy-3-methylphenyl, 3-methylaminophenyl, 4-methylaminophenyl, 4-ethylaminophenyl or 2-aminomethylphenyl;

(ii) naphth-2-yl, 3-aminonaphth-2-yl, 3-hydroxynaphth-2-yl or 6-hydroxynaphth-2-yl;

(iii) isoquinolin-7-yl, indol-5-yl, indol-6-yl, 3-chloroindol-6-yl, 3-bromoindol-6-yl, 3-methylindol-6-yl, 3-methoxyindol-6-yl, indazol-5-yl, 3-aminoindazol-5-yl, indazol-6-yl, benzothiazol-6-yl, 3-aminobenzisoxazol-5-yl;

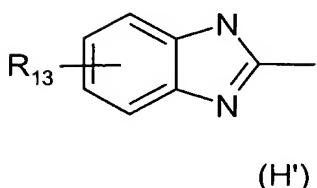
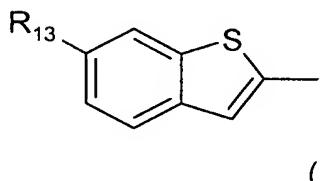
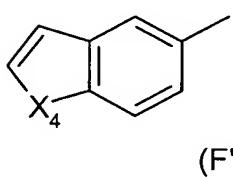
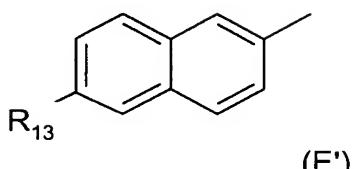
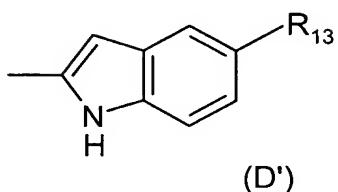
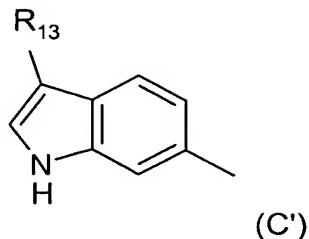
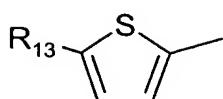
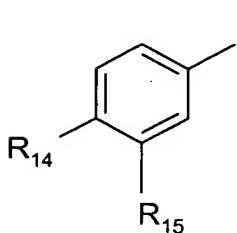
(iv) benzimidazol-5-yl, 2-aminobenzimidazol-5-yl, or benzothiazol-6-yl;

(v) thien-2-yl, 5-methylthien-2-yl, 5-methylthio-thien-2-yl, 5-acetylthien-2-yl or thien-3-yl;

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- (vi) 3,4-methylenedioxypyphenyl, 2,3-dihydroindol-6-yl,  
3,3-dichloro-2-oxo-indol-6-yl or 1-methyl-3-aminoindazol-5-yl;
- (vii) benzothiazol-2-yl, imidazo[1,2-a]pyrimidin-2-yl or  
tetrahydroimidazo[1,2-a]pyrimidin-2-yl;
- (viii) 5-methylpyrazol-2-yl;
- (ix) 5-chloropyrid-2-yl;
- (x) pyrid-3-yl, 6-chloropyrid-3-yl;
- (xi) benzofur-2-yl, 5-chlorobenzofur-2-yl, 3-  
methylbenzofur-2-yl, 5-methylbenzofur-2-yl, 6-methoxybenzofur-  
2-yl;
- (xii) indol-2-yl, 5-fluoroindol-2-yl, 5-chloroindol-2-yl,  
5-methylindol-2-yl, 5-methoxindol-2-yl, 6-methoxyindol-2-yl  
and 1-methyl-indol-2-yl;
- (xiii) 5-fluoroindol-6-yl; or
- (xiv) benzo[b]thiophen-2-yl, 5-chloro- benzo[b]thiophen-  
2-yl or 6-chlorobenzo[b]thiophen-2-yl. ~~fluoro, chlоро, bromо,~~  
~~iodо, nitro, thiol, difluoromethoxy, trifluoromethoxy,~~  
~~hydrazido, methylhydrazido, amino, cyano, trifluoromethyl,~~  
~~methylthio, vinyl, ethynyl, acetylarnino, carboxy, acetoxу,~~  
~~hydroxy, methyl, ethyl, amide (CONH<sub>2</sub>), aminomethyl, methoxy~~  
~~and ethoxy.~~

11 (currently amended): A compound according to claim any one  
of claims 1 to 10 wherein R<sub>2</sub> is selected from one of the  
formula (A') to (H'):



wherein  $X_4$  is O or S,  $R_{13}$  is selected from hydrogen, fluoroo-, [except for (C')], chloro or methyl and  $R_{14}$  is selected from hydrogen, methyl, ethyl, fluoro, chloro, and methoxy and  $R_{15}$  is selected from hydrogen, methyl, fluoro, chloro and amino.

12 (currently amended): A compound according to claim claims 1-to-11, wherein  $R_2$  is 4-chlorophenyl, 4-methoxyphenyl, 3-amino-4-chlorophenyl, indol-2-yl, 5-chloroindol-2-yl, indol-6-yl, 3-chloroindol-6-yl or 3-methylindol-6-yl.

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13 (canceled) :

14 (canceled) :

15 (canceled) :

16 (canceled) :

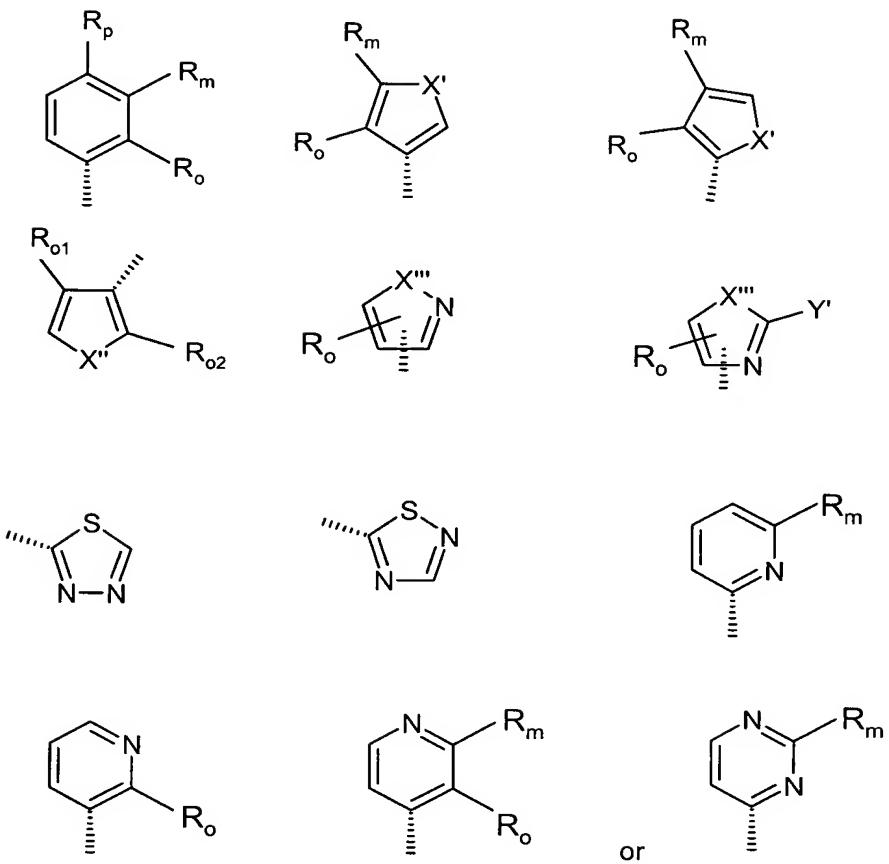
17 (canceled) :

18 (canceled) :

19 (currently amended) : A compound according to claim 1 any one of claims 1 to 16 wherein R<sub>3a</sub> is selected from hydrogen, hydroxyl, methoxy, ethoxy, methyl, ethyl, methylaminomethyl, dimethylaminomethyl, hydroxymethyl, carboxy, methoxymethyl, methoxycarbonyl, ethoxycarbonyl, methylaminocarbonyl, dimethylamino-carbonyl, aminomethyl, CONH<sub>2</sub>, CH<sub>2</sub>CONH<sub>2</sub>, acetylarnino, methoxycarbonylamino, ethoxycarbonylamino, t-butoxycarbonylamino, amino, fluoro, chloro, bromo, cyano, nitro, thiol, methylthio, methylsulphonyl, ethylsulphonyl, methylsulphenyl, methylsulphonylamido, ethylsulphonylamido, methylaminosulphonyl, ethylaminosulphonyl, aminosulphonyl, trifluoromethoxy, trifluoromethyl, bromo, -OCH<sub>2</sub>O- (which is bonded to two adjacent ring atoms in Cy) and -C(X<sup>3</sup>)N(R<sup>11</sup>)R<sup>12</sup> (wherein X<sup>3</sup> is O or S and R<sup>11</sup> and R<sup>12</sup> are independently selected from hydrogen, methyl or ethyl or together with the nitrogen atom to which they are attached form a pyrrolidin-1-yl, piperidin-1-yl or morpholino group).

20 (canceled) :

21 (currently amended) : A compound according to claim any one of claims 1 to 14 wherein Cy is selected from:



wherein:

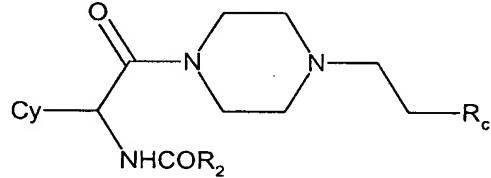
X' is selected from O, S and NMe;  
 X'' is selected from O and S;  
 X''' is selected from O, S, NH and NMe;  
 Y' is selected from hydrogen, amino and methyl;  
 R<sub>o</sub> is selected from hydrogen, methyl, fluoro, chloro, trifluoromethyl, methoxy, methylthio, methylsulphanyl and methylsulphonyl;  
 R<sub>m</sub> is selected from hydrogen, methyl, fluoro, chloro, trifluoromethyl, methoxy, methylthio, methylsulphanyl, methylsulphonyl, carboxy, methoxycarbonyl and a group of the

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formula  $-C(X^3)N(R^{11})R^{12}$  (wherein  $X^3$  is O or S and  $R^{11}$  and  $R^{12}$  are independently selected from hydrogen, methyl or ethyl or together with the nitrogen atom to which they are attached form a pyrrolidin-1-yl, piperidin-1-yl or morpholino group);  $R_p$  is selected from hydrogen and fluoro; or  $R_O$  and  $R_m$  or  $R_m$  and  $R_p$  form an  $-OCH_2O-$  group; or  $R_O$  and  $R_m$  together with the ring to which they are attached form a 5 or 6 membered aryl or heteroaryl ring (wherein the heteroary ring contains 1 or 2 heteroatoms selected from nitrogen, oxygen and sulfur); and one of  $R_{O1}$  and  $R_{O2}$  is hydrogen and the other is  $R_{O\frac{1}{2}}$ .

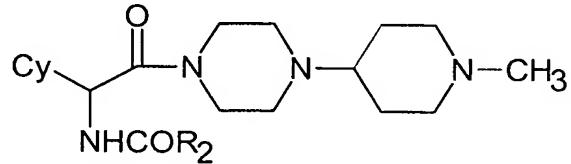
22 (currently amended): A compound according to claim any one of claims 1 to 14 wherein Cy is selected from phenyl, 2-chlorophenyl, 2-methoxyphenyl, 4-carbamoylphenyl, pyrid-2-yl, pyrid-4-yl, thien-2-yl, thien-3-yl, furan-2-yl, furan-3-yl, imidazol-2-yl, thiazol-2-yl, thiazol-4-yl, thiazol-5-yl and quinolin-4-yl.

23 (currently amended): A compound of the formula:



wherein Cy, R<sub>2</sub> and R<sub>c</sub> are as defined in any one of claims 1 to 226, 9 to 12, 19 and 21 to 22.

24 (currently amended): A compound of the formula:



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wherein Cy and R<sub>2</sub> are as defined in any one of claims 1 to  
222, 9 to 12, 19 and 21 to 22.

25 (canceled) :

26 (currently amended) : A compound as claimed in Claim 1,  
which is selected from:

1-(Indole-6-carbonyl-D-phenylglycinyl)-4-[2-(4-pyridinyl)-ethyl]piperazine;  
1-(3-Chloroindole-6-carbonyl-D-phenylglycinyl)-4-[2-(4-pyridinyl)ethyl]piperazine;  
1-(4-Methoxybenzoyl-D-phenylglycinyl)-4-(1-methylpiperidin-4-yl)piperazine;  
~~1-(Indole-6-carbonyl-D-phenylglycinyl)-4-(1-methylpiperidin-4-yl)piperazine;~~  
1-(4-Methoxybenzoyl-D-(2-chlorophenyl)glycinyl)-4-(1-methylpiperidin-4-yl)piperazine;  
1-(Indole-6-carbonyl-D-(2-chlorophenyl)glycinyl)-4-(1-methylpiperidin-4-yl)piperazine; and  
1-(4-Methoxybenzoyl-D-(2-trifluoromethylphenyl)glycinyl)-4-(1-methylpiperidin-4-yl)piperazine;  
and physiologically-tolerable salts thereof.

27 (currently amended) : A pharmaceutical composition, which comprises a compound as claimed in any one of Claims 1 to 26 together with at least one pharmaceutically acceptable carrier or excipient.

28 (canceled) :

29 (canceled) :

30 (currently amended) : A method of treatment of a human or non-human animal body to combat a thrombotic disorder selected

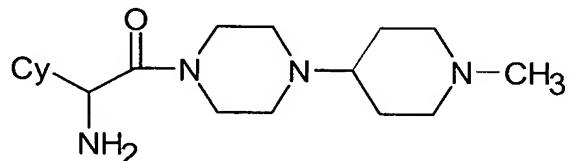
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from venous thrombosis, pulmonary embolism, arterial thrombosis, myocardial ischaemia, myocardial infarction and cerebral thrombosis, which comprises administering to said body an effective amount of a compound as claimed in claim 1.

31 (canceled) :

32 (canceled) :

33 (currently amended) : A compound of the formula



or a salt thereof in which Cy is as defined in any one of claims 1, 21 and 22.

34 (new) : A compound as claimed in any one of claims 1 to 13, 19 and 21 to 22, wherein the alpha atom in Y is carbon and has the conformation that would result from construction from a D- $\alpha$ -aminoacid NH<sub>2</sub>-CH(Cy)-COOH where the NH<sub>2</sub> represents part of X-X.

35 (new) : A pharmaceutical composition, which comprises a compound as claimed in claim 34 together with at least one pharmaceutically acceptable carrier or excipient.

36 (new) : A method of treatment of a human or non-human animal body to combat a thrombotic disorder selected from venous thrombosis, pulmonary embolism, arterial thrombosis, myocardial ischaemia, myocardial infarction and cerebral thrombosis, which comprises administering to said body an effective amount of a compound as claimed in claim 34.

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37 (new) : A method as claimed in claim 36 in which said human or non-human animal body is a human body.